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For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

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(54) Title: COMPOSITIONS AND METHODS FOR DETECTION AND TREATMENT OF PROLIFERATIVE ABNORMALITIES ASSOCIATED WITH OVEREXPRESSION OF HUMAN TRANSKETOLASE LIKE-1 GENE

(57) Abstract: The present invention relates to methods for treatment and diagnosis of disorders associated with abnormally proliferating cells. In one aspect the invention relates to methods, which are especially useful for the detection of tumors and their precursory stages based on the detection of overexpression of human transketolase like-1 gene in biological samples. In another aspect the invention relates to methods for treatment of disorders associated with the overexpression of human transketolase like-1 gene. Methods for treatment may include gene therapeutic approaches as well as methods for inhibiting or reducing the activity of transketolase like-1 polypeptides.

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PCT/EP 03/03827

	_	PCIA	/EP 03/0382/						
A. CLASSII IPC 7	FICATION OF SUBJECT MATTER C12Q1/68								
According to	International Patent Classification (IPC) or to both national classific	ation and IPC							
B. FIELDS	SEARCHED								
Minimum do IPC 7	cumentation searched (classification system followed by classification ${\tt C12Q}$	on symbols)							
Documentat	ion searched other than minimum documentation to the extent that a	such documents are included. In t	he fields searched						
Electronic da	ata base consulted during the International search (name of data ba	se and, where practical, search	terms used)						
EPO-Internal, BIOSIS, PAJ, WPI Data									
C. DOCUME	INTS CONSIDERED TO BE RELEVANT								
Category °	Citation of document, with indication, where appropriate, of the rel	Relevant to claim No.							
х	COY J F ET AL: "MOLECULAR CLONINTISSUE-SPECIFIC TRANSCRIPTS OF A	18-20							
	TRANSKETOLASE-RELATED GENE: IMPLIFOR THE EVOLUTION OF NEW VERTEBRA								
	GENOMICS, ACADEMIC PRESS, SAN DIE	:GU, US,							
	15 March 1996 (1996-03-15), pages XP000616489								
Y	ISSN: 0888-7543 the whole document	1–29							
χ	WO 97 05253 A (DEUTSCHES KREBSFOF ;POUSTKA ANNEMARIE (DE); COY JOHA	18-20							
Υ	(DE)) 13 February 1997 (1997-02-1 the whole document	.3)	1-29						
	-	-/							
X Furth	er documents are listed in the continuation of box C.	X Patent family members	e are listed in annex.						
Special cat	tegories of cited documents :	*T* later document published aff	er the International filing date						
"A" docume conside "E" earlier d	onflict with the application but ciple or theory underlying the ance; the claimed invention								
*L* document which is	of cannot be considered to then the document is taken alone ance; the claimed invention								
*O* docume	or other special reason (as specified) ent referring to an oral disclosure, use, exhibition or	volve an inventive step when the none or more other such docu-							
other n "P" docume later th	eing obvious to a person skilled me patent family								
Date of the a	Date of the actual completion of the international search  Date of mailing of the international search report								
8	July 2003	15/07/2003	15/07/2003						
Name and m	naiting address of the ISA	Authorized officer							
	European Patent Office, P.B. 5818 Patentiaan 2 NL - 2280 HV Rijswijk								
	Tel. (+31-70) 340-2040. Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Bort, S							

Int. anal Application No
PCT/EP 03/03827

Category Citation of document, with indication, where appropriate, of the relevant passages  X CASCANTE ET AL.: "Role of thiamin (vitamin B-1) and transketolase in tumor cell proliferation" NUTRITION AND CANCER, vol. 36, no. 2, 2000, pages 150-154, XP008012308 the whole document	Relevant to claim No.  30,33
X CASCANTE ET AL.: "Role of thiamin (vitamin B-1) and transketolase in tumor cell proliferation" NUTRITION AND CANCER, vol. 36, no. 2, 2000, pages 150-154, XP008012308 the whole document	30,33
(vitamin B-1) and transketolase in tumor cell proliferation" NUTRITION AND CANCER, vol. 36, no. 2, 2000, pages 150-154, XP008012308 the whole document	
W 000000000000000000000000000000000000	31,32
X SPENTZOS ET AL.: "Transketolase gene expression may be a marker of sensitivity to platinum/paclitaxel chemotherapy in ovarian cancer" PROCEEDINGS OF THE AMERICAN ASSOCIATION FOR CANCER RESEARCH ANNUAL MEETING, vol. 43, March 2002 (2002-03), pages 1105-1106, XP008012309	
Y the whole document	1-29
LYNCH ET AL.: "Hereditary cancer: family history, diagnosis, molecular genetics, ecogenetics, and management strategies" BIOCHIMIE, vol. 84, no. 1, January 2002 (2002-01), pages 3-17, XP001148125 the whole document	1-17
WO 98 53319 A (KINZLER KENNETH W ; VOGELSTEIN BERT (US); UNIV JOHNS HOPKINS (US)) 26 November 1998 (1998-11-26) abstract; claims 2,29,52	1-17
WO 00 52204 A (ORNTOFT TORBEN F) 8 September 2000 (2000-09-08) abstract; claims 8,12,22	1-17

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Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This international Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. X Claims Nos.: 21-27, 30-33 (all partially) because they relate to subject matter not required to be searched by this Authority, namely:
see FURTHER INFORMATION sheet PCT/ISA/210
2. X Claims Nos.: 30, 33 (all partially) because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
see FURTHER INFORMATION sheet PCT/ISA/210
a Chilma Nasa
Claims Nos.:  because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This international Searching Authority found multiple inventions in this international application, as follows:
As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. As all searchable dalms could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the Invention first mentioned in the claims; it is covered by claims Nos.;
Remark on Protest  The additional search fees were accompanied by the applicant's protest.
No protest accompanied the payment of additional search fees.

International Application No. PCTEP 03 03827

#### FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box I.1

Although claims 21-27, 31 and 32 are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effect.

Continuation of Box I.1

Claims Nos.: 21-27, 30-33 (all partially)

Rule 39.1(iv) PCT - Method for treatment of the human or animal body by therapy

Continuation of Box I.2

Claims Nos.: 30, 33 (all partially)

Independent claim 30 refers to a pharmaceutical compositions for the treatment of certain types of cancer, defined by reference to a desirable characteristic or property, namely the method fo claim 29. The claim covers all compositions having this characteristic although the application does not provide support therefor within the meaning of Article 6 PCT. The lack of support/disclosure in the present case is such that any compound used to treat cancer will fall under the scope of protection. Consequently, the search of claim 30 has been limited to any antihiamine compound, inhibitor or transketolase anzyme activity, transketolase like-1 polypeptide or human transketolase like-1 nucleic acid

The applicant's attention is drawn to the fact that claims, or parts of claims, relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure.

information on patent family members

Interest PCT/EP 03/03827

Patent document cited in search report		Publication date	Patent family member(s)		Publication date
WO 9705253	A	13-02-1997	DE WO EP JP US	19527552 A1 9705253 A2 0840789 A2 11510382 T 6133005 A	30-01-1997 13-02-1997 13-05-1998 14-09-1999 17-10-2000
WO 9853319	A	26-11-1998	AU WO	7499198 A 9853319 A2	11-12-1998 26-11-1998
WO 0052204	Α	08-09-2000	AU WO US	3316600 A 0052204 A2 6335170 B1	21-09-2000 08-09-2000 01-01-2002